

(FILE 'HOME' ENTERED AT 08:05:24 ON 19 MAR 1999)

FILE 'USPATFULL' ENTERED AT 08:05:32 ON 19 MAR 1999

FILE 'MEDLINE, EMBASE, CAPLUS' ENTERED AT 08:05:53 ON 19 MAR 1999  
1276 S MESODERM? AND (BMP? OR HEDGEHOG OR WNT? OR VEGF?)

L1  
L2 97 S L1 AND (HEMATOP? OR HEMATAO? OR HEAMATOP? OR VASCUL?)

FILE 'MEDLINE' ENTERED AT 08:16:10 ON 19 MAR 1999

FILE 'MEDLINE, EMBASE, CAPLUS' ENTERED AT 08:19:19 ON 19 MAR 1999

L3 52 S L2 NOT PY>1997  
L4 24 DUP REM L3 (28 DUPLICATES REMOVED)  
E BARON M/AU  
L5 129 S E9  
E FARRINGTON S/AU  
L6 25 S E5-E7  
E BARON MARG/AU  
L7 26 S E5  
E BELAOUSSOFF/AU  
L8 9 S E4-E5  
L9 170 S L5 OR L6 OR L7 OR L8  
E HEMATO?/  
L10 9 S L9 AND (HEMATOP? OR HEAMATOP? OR HEMATAO? OR VASCUL?)  
L11 7 DUP REM L10 (2 DUPLICATES REMOVED)

(FILE 'USPAT' ENTERED AT 15:25:13 ON 18 MAR 1999)

L1 1013 S BMP? OR HEDGEHOG OR WNT?  
E HEMAT/  
E HEAMAT/  
L2 3289 S E14 OR E15 OR HEMATAOPO? OR HEMATOPO?  
L3 103 S L1 AND L2  
L4 3 S L1 (15A) (VASCUL?)  
L5 77 S L3 AND (EMBRYO? OR MESODERM? OR UNDIFFERENT?)

Novel

*Vascular  
endothelium from mesoderm*

*11*  
*12*  
*13*  
*14*  
*15*  
*16*  
*17*  
*18*  
*19*  
*20*  
*21*  
*22*

L4 ANSWER 19 OF 24 LINE PLICATE 12  
TI The angiogenic potentials of the cephalic **mesoderm** and the  
origin of brain and head blood vessels.  
AU Couly G; Coltey P; Eichmann A; Le Douarin N M  
SO MECHANISMS OF DEVELOPMENT, (1995 Sep) 53 (1) 97-112.  
Journal code: AXF. ISSN: 0925-4773.  
AB We have used two molecular markers to label blood vessel endothelial  
cells  
and their precursors in the early avian embryo. One marker, called Quek1,  
is the avian homologue of the mammalian **VEGF** receptor flk-1 and  
the other is the MB1/QH1 monoclonal antibody. Quek1 is expressed in a  
subset of **mesodermal** cells from the gastrulation stage. Quek1  
positive cells later form blood vessel endothelial cells and express the  
MB1/QH1 antigen which is specific for endothelial and hemopoietic cells  
of  
the quail species. These two markers allowed us first to show that the  
cephalic paraxial **mesoderm** has angiogenic potentials which are  
much more extended than its trunk counterpart (the somites). Secondly,  
the  
origin of the endothelial cells lining the craniofacial and head blood  
vessels was mapped on the 3-somite stage cephalic **mesoderm** via  
the quail-chick chimera technique, in which well defined  
**mesodermal** territories are exchanged between stage-matched embryos  
of both species in a strictly isotopic manner. We found that the anterior  
region of the cephalic paraxial **mesoderm** is largely recruited to  
provide the forebrain and the upper face with their **vasculature**.  
This means that large volumes of tissues are **vascularized** by a  
discrete region of the cephalic **mesoderm**, the fate of which is  
otherwise to give rise to muscles. The widespread expansion of the  
angiogenic cells arising from the anterior paraxial **mesoderm**  
must be related to the high growth rate of the anterior region of the  
neural primordium, yielding the telencephalon and of the neural  
crest-derived facial structures which are themselves devoid of angiogenic

TI Development and differentiation of blood vessels in the central nervous system

AU Wilting, J.

SO Neuroendocrinol., [Ernst Berta Scharrer Symp.] (1997), 305-312.

Editor(s): Korf, Horst-Werner; Usadel, Klaus-Henning. Publisher:

Springer,

Berlin, Germany.

CODEN: 66EOAA

AB A review, with 44 refs. The central nervous system (CNS) develops from a pseudostratified ectodermal epithelium contg. neuroblasts and glioblasts. Other constituents (microglia, blood vessels) are of **mesodermal** origin and successively invade the neuroectoderm. Using chick-quail chimeras it is possible to study the interaction between neuroectodermal and **mesodermal** cells. **Vascular** endothelial cells start invading the CNS of birds at about day 3.5 of development. They originate from the paraxial **mesoderm** of the head and the trunk. Thereafter, smooth muscle cells migrate along the endothelial routes. Neuroectodermal cells secrete **vascular** endothelial growth factor (**VEGF**), which is a highly specific angiogenic and chemoattractive factor. Angioblast and endothelial cells in the paraxial **mesoderm** are characterized by the expression of **VEGF**-receptor-2. Except for the choroid plexus, **VEGF** and **VEGF** receptors are not expressed in the adult brain. The organ-typical differentiation of endothelial cells in the CNS depends on interactions with local neuroectodermal cells. Development of

blood-brain

barrier characteristics are obviously due to inductive signals from astrocytes. In contrast, the epithelial cells of the choroid plexus induce development of highly permeable, fenestrated capillaries. Constitutive expression of **VEGF** and its receptors in the choroid plexus (and the kidney glomeruli) may serve as the basis for high permeability. **VEGF** has been shown to increase **vascular** permea

- TI **Vascularization** of the mouse embryo: a study of flk-1, tek, tie, and **vascular** endothelial growth factor expression during development.
- AU Dumont D J; Fong G H; Puri M C; Gradwohl G; Alitalo K; Breitman M L
- SO DEVELOPMENTAL DYNAMICS, (1995 May) 203 (1) 80-92.  
Journal code: A9U. ISSN: 1058-8388.
- AB We report the detailed developmental expression profiles of three endothelial specific receptor tyrosine kinases (RTKs) flk-1, tek, tie, as well as **vascular** endothelial growth factor (**VEGF**), the flk-1 ligand. We also examined the expression of the other **VEGF** receptor, flt-1, during placental development. flk-1, tek, and tie transcripts were detected sequentially at one-half day intervals starting at E7.0, suggesting that each of these RTKs play a unique role during **vascularization** of the mouse embryo. All three RTKs were expressed in the extraembryonic and embryonic **mesoderm** in regions that eventually give rise to the **vasculature**. Except for the expression of tek and flk-1 in the **mesoderm** of the amnion, the expression of these RTKs from E8.5 onwards was virtually indistinguishable. An abundant amount of flt-1 transcripts was found in the spongiotrophoblast cells of the developing placenta from E8.0 onwards.
- This cellular compartment is located between the maternal and labyrinthine layers of the placenta, which both express **VEGF**. **VEGF** transcripts were detected as early as E7.0 in the endoderm juxtaposed to the flk-1 positive **mesoderm**, and later in development **VEGF** expression displayed an expression profile both contiguous with that of flk-1, and also in tissues found some distance from the flk-1-expressing endothelium. These results suggest a possible dual role for **VEGF** which includes a chemotactic and/or a cellular maintenance role for **VEGF** during **vascularization** of the mouse emb

L4 ANSWER 14 OF 24 MEDLINE

DUPLICATE 8

TI In vitro analysis of epiblast tissue potency for **hematopoietic** cell differentiation.

AU Kanatsu M; Nishikawa S I

SO DEVELOPMENT, (1996 Mar) 122 (3) 823-30.

Journal code: ECW. ISSN: 0950-1991.

AB In murine embryogenesis, all cells that will constitute the embryonic structures originate from the epiblast (primitive ectoderm) tissue, the epithelial cell sheet of the gastrulating embryo. The cells of this tissue

are totipotent at the beginning of gastrulation, but at the end of this period are specified to particular cell lineages. Thus, it is likely that during murine gastrulation, the potency of epiblast cells that were originally totipotent becomes restricted as development progresses. However, the mechanisms of this process are unknown. We have investigated this process in vitro, focusing on the **hematopoietic** cell lineage. To detect the hematogenic potency of the epiblast tissue, we established an in vitro culture system in which the **hematopoietic** cell differentiation of the epiblast tissue was supported by a stromal cell layer. With this culture system, we investigated the process by

which

this potency becomes spatially and temporally restricted during gastrulation. The results showed that hematogenic potency resides in the entire epiblast of the early- to mid-gastrulating embryo, but becomes restricted to the posterior half of the epiblast at the headfold stage. Furthermore, we showed that this process is altered by exogenous bone morphogenetic protein-4 (**BMP-4**) or activin A, which may be **mesoderm** inducers in *Xenopus* embryogenesis.

- L11 ANSWER 1 OF 7 CAPLUS COPYRIGHT 1999 ACS  
 TI Embryonic induction of **hematopoietic** and **vascular**  
 mesoderm in the developing mouse  
 AU **Belaoussoff, Maria**  
 SO (1998) 211 pp. Avail.: UMI, Order No. DA9832323  
 From: Diss. Abstr. Int., B 1998, 59(5), 2017
- L11 ANSWER 2 OF 7 CAPLUS COPYRIGHT 1999 ACS  
 TI Methods for modulating **hematopoiesis** and **vascular**  
 growth  
 IN **Baron, Margaret H.; Farrington, Sarah M.;**  
**Belaoussoff, Maria**  
 SO PCT Int. Appl., 77 pp.  
 CODEN: PIXXD2
- L11 ANSWER 3 OF 7 MEDLINE DUPLICATE 1  
 TI **Hematopoietic** induction and respecification of A-P identity by  
 visceral endoderm signaling in the mouse embryo.  
 AU **Belaoussoff M; Farrington S M; Baron M H**  
 SO DEVELOPMENT, (1998 Dec) 125 (24) 5009-18.  
 Journal code: ECW. ISSN: 0950-1991.
- L11 ANSWER 4 OF 7 CAPLUS COPYRIGHT 1999 ACS  
 TI A novel developmental regulatory motif required for stage-specific  
 activation of the .epsilon.-globin gene and nuclear factor binding in  
 embryonic erythroid cells  
 AU Trepicchio, William L.; Dyer, Michael A.; **Baron, Margaret H.**  
 SO Mol. Cell. Biol. (1994), 14(6), 3763-71  
 CODEN: MCEBD4; ISSN: 0270-7306
- L11 ANSWER 5 OF 7 CAPLUS COPYRIGHT 1999 ACS  
 TI Positive regulators of the lineage-specific transcription factor GATA-1  
 in  
 differentiating erythroid cells  
 AU **Baron, Margaret H.; Farrington, Sarah M.**  
 SO Mol. Cell. Biol. (1994), 14(5), 3108-14  
 CODEN: MCEBD4; ISSN: 0270-7306
- L11 ANSWER 6 OF 7 CAPLUS COPYRIGHT 1999 ACS  
 TI Developmental regulation of the human embryonic .beta.-like globin gene  
 is  
 mediated by synergistic interactions among multiple tissue- and  
 stage-specific elements  
 AU Trepicchio, William L.; Dyer, Michael A.; **Baron, Margaret H.**  
 SO Mol. Cell. Biol. (1993), 13(12), 7457-68  
 CODEN: MCEBD4; ISSN: 0270-7306
- L11 ANSWER 7 OF 7 CAPLUS COPYRIGHT 1999 ACS  
 TI Reprogramming of globin gene expression in interspecific heterokaryons  
 AU **Baron, Margaret H.; Maniatis, Tom**  
 SO UCLA Symp. Mol. Cell. Biol., New Ser. (1987), 51(Mol. Approaches Dev.  
 Biol.), 469-76  
 CODEN: USMBD6; ISSN: 0735-9543

L1 ANSWER 1 OF 2 INPADOC COPYRIGHT 1999 EPO  
 AN 28952038 INPADOC UW 9909 UP 990313 EW 9909 ED 990313  
 TI METHODS FOR MODULATING HEMATOPOIESIS AND VASCULAR GROWTH.  
 IN BARON, MARGARET, H.; FARRINGTON, SARAH, M.; BELAOUSOFF, MARIA  
 INS BARON MARGARET H; FARRINGTON SARAH M; BELAOUSOFF MARIA  
 PA THE PRESIDENTS AND FELLOWS OF HARVARD COLLEGE  
 PAS PRESIDENTS AND FELLOWS OF HARV  
 PAA US  
 LA English  
 TL English; French  
 DT Patent  
 PIT WOA3 SUBSEQUENT PUBL. OF THE INT. SEARCH REPORT  
 PI WO 9835020 A3 990114 300000  
 DS W CA; W JP  
 RW AT; RW BE; RW CH; RW DE; RW DK; RW ES; RW FI; RW FR; RW GB; RW GR; RW  
 IE; RW IT; RW LU; RW MC; RW NL; RW PT; RW SE  
 AI WO 98-US2633 A 980210  
 PRAI US 97-37513 P 970210 EWPR 9835 EDPR 980905  
 US 97-49763 P 970616 EWPR 9835 EDPR 980905

VEGF + TLE TEK

45,585,087

5,585,237

5,565,321